

ABSTRACT OF THE DISCLOSURE

The present invention relates to a transgenic non-human mammal whose germ cells and somatic cells contain a knockout mutation in DNA encoding orphan nuclear receptor $ERR\alpha$. More specifically, the invention relates to

5 knockout mice whose germ cells and somatic cells contain a knockout mutation in DNA encoding orphan nuclear receptor $ERR\alpha$. The invention further relates to such knockout non-human mammals which express an $ERR\alpha$ gene which is different from the endogenous gene which was disrupted. In a particular embodiment, the invention relates to a transgenic mouse having its endogenous

10 $ERR\alpha$ gene disrupted and expressing human $ERR\alpha$. As well, the invention relates to cell lines in which $ERR\alpha$ activity (and/or level) has been inactivated or augmented. The invention further relates to uses and methods of the transgenic animals of the present invention to select agents which modulate the expression and/or activity of $ERR\alpha$.